

ation No International A PCT/EP 03/03060

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 A61K31/44 A61K31/45 A61P35/00 A61P43/00

A61K31/495

A61K31/4406 A61P37/00

According to International Patent Classification (IPC) or to both national classification and IPC

Minimum documentation searched (classification system followed by classification symbols)  $IPC\ 7\ A61K$ 

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

	ternal, WPI Data, PAJ, CHEM ABS Dat	u, p10010		
C. DOCUME	ENTS CONSIDERED TO BE RELEVANT			
Category °	Citation of document, with indication, where appropriate, of the re	elevant passages	Relevant to claim No.	
х	DE 196 24 659 A (KLINGE CO CHEM 8 January 1998 (1998-01-08) page 64; examples 150,153,158,15 page 36 page 2, line 1-26 page 77, line 51 -page 78, line	9,162	1-11,14	
X	DE 197 56 235 A (KLINGE CO CHEM 1 July 1999 (1999-07-01) page 33; examples 82,120 page 74, line 40 -page 75, line		1-11,14	
X	DE 196 24 704 A (KLINGE CO CHEM 8 January 1998 (1998-01-08) claim 9 page 78 -page 79, line 7	PHARM FAB)	1-11,14	
X Furl	her documents are listed in the continuation of box C.	-/  γ Patent family members are listed	in annex.	
Special ca     A docume consider     E earlier filing of the charto charto     O docume other     P docume     P docume	ent defining the general state of the art which is not letted to be of particular relevance document but published on or after the international late ent which may throw doubts on priority claim(s) or is cited to establish the publication date of another n or other special reason (as specified) ent referring to an oral disclosure, use, exhibition or means ent published prior to the international filing date but han the priority date claimed	<ul> <li>"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention</li> <li>"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone</li> <li>"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.</li> <li>"&amp;" document member of the same patent family</li> </ul>		
Date of the	actual completion of the International search	Date of mailing of the international se	earch report	
9	July 2003	21/07/2003		
Name and	malling address of the ISA  European Patent Office, P.B. 5818 Patentlaan 2  NL - 2280 HV Rijswijk  Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Authorized officer  Bonzano, C		



International A action No PCT/EP 03/03060

C.(Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	DE 197 56 212 A (KLINGE CO CHEM PHARM FAB) 1 July 1999 (1999-07-01)	
A	DE 197 56 236 A (KLINGE CO CHEM PHARM FAB) 1 July 1999 (1999-07-01)	
Υ	US 4 778 796 A (UNO HITOSHI ET AL) 18 October 1988 (1988-10-18) column 24; example 85 column 25; example 91 column 1, line 7 - line 37	1-11
X -	DE 197 56 261 A (KLINGE CO CHEM PHARM FAB) 1 July 1999 (1999-07-01) page 2, line 21 - line 50 page 38	1-6,11
<b>Y</b>	NIE DAOTAI ET AL: "Eicosanoid regulation of angiogenesis: Role of endothelial arachidonate 12-lipoxygenase." BLOOD, vol. 95, no. 7, 1 April 2000 (2000-04-01), pages 2304-2311, XP002208812 ISSN: 0006-4971 page 2310, column 1, paragraph 1 page 2310, column 2, paragraph 3 - paragraph 4	1-11
Α	COLAVITTI RENATA ET AL: "Reactive oxygen species as downstream mediators of angiogenic signaling by vascular endothelial growth factor receptor-2/KDR." JOURNAL OF BIOLOGICAL CHEMISTRY, vol. 277, no. 5, February 2002 (2002-02), pages 3101-3108, XP001074657 February, 2002 ISSN: 0021-9258 page 3101, column 2, paragraph 3	
Υ	NISHIKAWA ET AL: "Acrylamide derivatives as antiallergic agents. 2. Synthesis and structure activity relationships of N-'4-'4-(diphenylmethyl)-1- piperazinyl!butyl!-3-(3-pyridyl)acrylamide s" JOURNAL OF MEDICINAL CHEMISTRY, AMERICAN CHEMICAL SOCIETY. WASHINGTON, US, vol. 32, no. 3, 1989, pages 583-593, XP002101517 ISSN: 0022-2623 page 587; examples 1A,16A page 588; examples 1A,21,1A; table V page 583, column 1, paragraph 1	1-11
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C.(Continua	ation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category °	Cliation of document, with Indication, where appropriate, of the relevant passages		Relevant to claim No.
A .	WO 98 02170 A (LEUVEN RESEARCH AND DEV V Z M ;GEN HOSPITAL CORP (US)) 22 January 1998 (1998-01-22) page 3, paragraph 2		
			·

## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

## Continuation of Box I.2

Claims 1-9,11,12 relate to the treatment of diseases and to diagnostic applications which are actually not well defined. The use of the definitions "inhibition or reduction of angiogenesis, disease responding to inhibition or reduction of angiogenesis, disease responding to inhibition or reduction of VEGF production; in vitro diagnostic method" in the present context is considered to lead to a lack of clarity within the meaning of Article 6 PCT. The lack of clarity is such as to render a meaningful complete search not fully possible.

Moreover, the claims cover all diseases responding to inhibition or reduction of angiogenesis and all in vitro diagnostic methods, whereas the application provides support within the meaning of Article 6 PCT and/or disclosure within the meaning of Article 5 PCT for only a very limited number of such diseases and diagnostic methods. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible.

Present claims 1-14 relate to an extremely large number of possible compounds. Support within the meaning of Article 6 PCT and disclosure within the meaning of Article 5 PCT is to be found, however, for only a very small proportion of the compounds claimed. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible. Consequently, the search has been restricted to the parts relating to the compounds mentioned in the examples, for the treatment of the diseases mentioned claims 10 and to the in vitro diagnostic methods for the diseases mentioned in claim 13, with due regard to the general idea underlying the present invention.

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

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## INTERNATIONAL SEARCH REPORT

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sh	eet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the followin	g reasons:
1. X Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:	
Although claim 14 is directed to a method of treatment of the human body, the search has been carried out and based on the alleged effe compound/composition.	/animal cts of the
2. X Claims Nos.:  Decause they relate to parts of the International Application that do not comply with the prescribed requirements to an extent that no meaningful International Search can be carried out, specifically:	o such
see FURTHER INFORMATION sheet PCT/ISA/210	
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Ru	le 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)	
This International Searching Authority found multiple inventions in this international application, as follows:	
As all required additional search fees were timely paid by the applicant, this international Search Report covers a searchable claims.	all
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite of any additional fee.	payment
3. As only some of the required additional search fees were timely paid by the applicant, this international Search for covers only those claims for which fees were paid, specifically claims Nos.:	Report
Concoulantly this International Search Re	enortis .
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Rerestricted to the invention first mentioned in the claims; it is covered by claims Nos.:	-e
Remark on Protest  The additional search tees were accompanied by the applic	ranta protest.
Remark on Protest  The additional search fees were accompanied by the application of additional search fees were accompanied by the additional search fees were accompanied by the application of additional search fees were accompanied by the application of additional search fees were accompanied by the application of additional search fees were accompanied by the additional search fees were accompanied by the additional search fees were accompanied by the additional search fees were accompani	
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